

PM International Studies
Series K Short Term Study (Osaka) – EHCK3K6/52/03/JP (MDS Study # TBD)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	<i>Aug 03</i>		<i>Aug 03</i>		
IRB Review	17 Oct 03	28 Oct 03			Needs to be completed by 17 Oct 03
Amendments Nr. 1					
Recruiting					
Screening					
Agreements/Proposals Execution Monitoring					
Study Conduct	Jan 04	Mar 04			5 sessions, 20 subjects
Topography Data					
Lab Analysis (clin and ext)					
SAP					
DB-Lock Clinical Analytical Statistical					
Preliminary Data Figs & Tabs					
Draft Report					
Monitoring					
Draft Report Review					
Final Report					
SDC/SDS					

Issues /Decisions	Resolution
1.	
2.	
3.	
4.	

Action items	Due Date	Done	MDS	PM
1.				
2.				
3.				
4.				

PM International Studies
Series K Long Term Study (Belfast)
EHCK3K6/51/03/UK (MDS Study # AA04861)

Approved
7/11/03

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol			04 Jul 03	31 Jul 03	
IRB Review	1 Aug 03	19 Aug 03			EC submission deadline 1 Aug 03
Amendments Nr. 1					
Recruiting					
Screening					
Agreements/Proposals Execution Monitoring					
Study Conduct	16 Oct 03 31 Oct 03	01 Apr 04 16 Apr 04			2 sessions
Topography Data					
Lab Analysis (clin and ext)					
SAP					
DB-Lock Clinical Analytical Statistical					
Preliminary Data Figs & Tabs					
Draft Report					
Monitoring					
Draft Report Review					
Final Report					
SDC/SDS					

Issues /Decisions	Resolution
1.	
2.	
3.	
4.	

Action items	Due Date	Done	MDS	PM
1.				
2.				
3.				
4.				

ACCORD SERIES K Short-term Study - EHCK6/01/03 (AA02981)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	-	-	23 Jan 03	30 Apr 03	
IRB Review	-	06 May 03	-	06 May 03	Approved (dated 01 May 03)
Amendments					
LOAC #1	-	-		27 May 03	IRB Approved (dated 22 May 03)
LOAC #2	-	-		17 Jun 03	IRB Approved (dated 17 Jun 03)
Nr1	06 Jun	10 Jun	06 Jun	10 Jun 03	Approved 10 Jun (dated 09 Jun 03)
Recruiting	29 Apr 03	2 Jun 03	29 Apr 03	02 Jun 03	470 subjects as of 4 Jun
Screening	3 Jun 03	12 Jun 03	03 Jun 03		June 3-6 screen, June 11-12 phys. Additional screening 11 Jun cancelled, phys 12 Jun
Agreements/Proposals					
Recruitment (PA)	-	-	14 Apr 03	29 Apr 03	MDS #42, Advertising, ICF
Screening, execution (PA)	15 May 03	02 Jun 03	16 May 03	02 Jun 03	MDS #44
44 Amend 1 for Prot Amend 1	-	21 Jul 03	18 Jul 03	30 Jul 03	To MDS 30 Jul 03—fully signed
Analysis-Scherer (PA)	14 May 03	29 Jun 03	19 May 03		#7, valine adducts
Analysis-Arista (AP/PO)	14 May 03	29 Jun 03	14 May 03	27 May 03	filter analysis
Analysis-Covance (PA)	14 May 03	29 Jun 03	28 May 03	30 Jul 03	#14, caffeine, 8-epi, 11-dehy
Monitoring (AP/PO)	14 May 03	29 Jun 03	25 May 03	08 Jul 03	Clinimetrics
Study Conduct	19 Jun 03	30 Jun 03	19 Jun 03	30 June 03	120 Subjects randomized and completed
Topography Data	30 Jun 03	15 Jul 03	30 Jun 03	15 Jul 03	
Lab Analysis (clin and ext)					
COHb	04 Jun 03	15 Jul 03	04 Jun 03	15 Jul 03	MDS
Caffeine Analysis				01 Aug 03	Covance
4-ABP	Needed	TBD			Covance
LDL, HDL, Hem, etc	Needed	TBD			MDS
Fib, Von, HSC, Alb	Needed	TBD			Esoterix/ UNMC
Valines	7 July 03	9 Aug 03	09 Jul 03	31 Jul 03	ABF (shipped 08 Jul)
Filter Analysis (N1409)	7 July 03	TBD			Arista (shipped 08 Jul) 1400 samples analyzed
SAP		21 Jul 03		27 Jun 03	ETS analysis 08/13/03
DB-Lock					
Clinical		18 Jul 03		21 Jul 03	
Analytical		08 Sep 03			Without MHBMA
Statistical		22 Sep 03			? soon
Preliminary Data					
Figs & Tabs		31 July 03			ETS ANAL → PD
Monitoring	22 July 03	15 Aug 03			22-25 July, 12-15 Aug, 19-22 Aug
Draft Report		30 Nov 03			
Draft Report Review	1 Dec 03	15 Dec 03			
Final Report		31 Dec 03			

Issues/Decisions	Resolution
1. Confirm whether MHBMA should remain in urinary biomarkers for study	YES as per a sec 60% 4/10/03 sent to covance

Action items	Due Date	Done	MDS	PM
1. Discussion around ETS analysis/process. Schedule teleconference. ETS data sent to Nancy week of 07/28/03.	23 Jul 03		KP	JO

ACCORD SERIES K Short-term Study - EHCK6/01/03 (AA02981)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	-	-	23 Jan 03	30 Apr 03	
IRB Review	-	06 May 03	-	06 May 03	Approved (dated 01 May 03)
Amendments					
LOAC #1	-	-		27 May 03	IRB Approved (dated 22 May 03)
LOAC #2				17 Jun 03	IRB Approved (dated 17 Jun 03)
Nr1	06 Jun	10 Jun	06 Jun	10 Jun 03	Approved 10 Jun (dated 09 Jun 03)
Recruiting	29 Apr 03	2 Jun 03	29 Apr 03	02 Jun 03	470 subjects as of 4 Jun
Screening	3 Jun 03	12 Jun 03	03 Jun 03		June 3-6 screen, June 11-12 phys. Additional screening 11 Jun cancelled, phys 12 Jun
Agreements/Proposals					
Recruitment (PA)	-	-	14 Apr 03	29 Apr 03	MDS #42, Advertising, ICF
Screening, execution (PA)	15 May 03	02 Jun 03	16 May 03	02 Jun 03	MDS #44
44 Amend 1 for Prot Amend 1	-	21 Jul 03	18 Jul 03	30 Jul 03	To MDS 30 Jul 03-fully signed
Analysis-Scherer (PA)	14 May 03	29 Jun 03	19 May 03	08 Aug 03	#7, valine adducts
Analysis-Arista (AP/PO)	14 May 03	29 Jun 03	14 May 03	27 May 03	filter analysis
Analysis-Covance (PA)	14 May 03	29 Jun 03	28 May 03	30 Jul 03	#14, caffeine, 8-epi, 11-dehy
Monitoring (AP/PO)	14 May 03	29 Jun 03	25 May 03	08 Jul 03	Clinimetrics
Study Conduct	19 Jun 03	30 Jun 03	19 Jun 03	30 June 03	120 Subjects randomized and completed
Topography Data	30 Jun 03	15 Jul 03	30 Jun 03	15 Jul 03	
Lab Analysis (clin and ext)					
COHB	04 Jun 03	15 Jul 03	04 Jun 03	15 Jul 03	MDS
Caffeine Analysis				01 Aug 03	Covance
4-ABP	<u>Needed</u>	<u>TBD</u>			Covance
LDL, HDL, Hem, etc	<u>Needed</u>	<u>TBD</u>			MDS
Fib, Von, HSC, Alb	<u>Needed</u>	<u>TBD</u>			Esoterix/ UNMC
Valines	7 July 03	31 Sep 03	09 Jul 03		ABF (shipped 08 Jul)
Filter Analysis	7 July 03	-	9 Jul 03	1 Aug 03	Arista, 1400 samples analyzed
SAP	21 Jul 03		27 Jun 03		Working on
DB-Lock					
Clinical		18 Jul 03		21 Jul 03	CCSR to QA wk of 25 Aug 03, need edit checks
Analytical		08 Sep 03			Without MHBMA (MHBMA
Statistical		22 Sep 03			locks 9/29 + 2 weeks for tables/figures)
Preliminary Data					
Figs & Tabs		31 July 03 13 Oct 03		31 Jul 03	COHB and topography tables Analytical tables and figures
Monitoring	22 July 03	<u>Mid Sep</u>			22-25 July, 12-15 Aug, 19-22 Aug, 26-29 Aug
Draft Report		1 Dec 03			
Draft Report Review	1 Dec 03	15 Dec 03			
Final Report		31 Dec 03			

Issues/Decisions	Resolution
1.	

Action items	Due Date	Done	MDS	PM
1. Summary tables for 3-HPMA data sent to PM as soon as possible	27 Aug 03	27 Aug 03	NW	MG
2.				

ACCORD SERIES K Short-term Study - EHCK6/01/03 (AA02981)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	-	-	23 Jan 03	30 Apr 03	
IRB Review	-	06 May 03	-	06 May 03	Approved (dated 01 May 03)
Amendments					
LOAC #1	-	-		27 May 03	IRB Approved (dated 22 May 03)
LOAC #2				17 Jun 03	IRB Approved (dated 17 Jun 03)
Nr1	06 Jun	10 Jun	06 Jun	10 Jun 03	Approved 10 Jun (dated 09 Jun 03)
Recruiting	29 Apr 03	2 Jun 03	29 Apr 03	02 Jun 03	470 subjects as of 4 Jun
Screening	3 Jun 03	12 Jun 03	03 Jun 03		June 3-6 screen, June 11-12 phys. Additional screening 11 Jun cancelled, phys 12 Jun
Agreements/Proposals					
Recruitment (PA)	-	-	14 Apr 03	29 Apr 03	MDS #42, Advertising, ICF
Screening, execution (PA)	15 May 03	02 Jun 03	16 May 03	02 Jun 03	MDS #44
44 Amend 1 for Prot Amend 1	-	21 Jul 03	18 Jul 03	30 Jul 03	To MDS 30 Jul 03—fully signed
Analysis-Scherer (PA)	14 May 03	29 Jun 03	19 May 03	08 Aug 03	#7, valine adducts
Analysis-Arista (AP/PO)	14 May 03	29 Jun 03	14 May 03	27 May 03	filter analysis
Analysis-Covance (PA)	14 May 03	29 Jun 03	28 May 03	30 Jul 03	#14, caffeine, 8-epi, 11-dehy
Monitoring (AP/PO)	14 May 03	29 Jun 03	25 May 03	08 Jul 03	Clinimetrics
Study Conduct	19 Jun 03	30 Jun 03	19 Jun 03	30 June 03	120 Subjects randomized and completed
Topography Data	30 Jun 03	15 Jul 03	30 Jun 03	15 Jul 03	
Lab Analysis (clin and ext)					
COHB	04 Jun 03	15 Jul 03	04 Jun 03	15 Jul 03	MDS
Caffeine Analysis				01 Aug 03	Covance
4-ABP	<u>Needed</u>	<u>TBD</u>			Covance
LDL, HDL, Hem, etc	<u>Needed</u>	<u>TBD</u>			MDS
Fib, Von, HSC, Alb	<u>Needed</u>	<u>TBD</u>			Esoterix/ UNMC
Valines	7 July 03	31 Sep 03	09 Jul 03		ABF (shipped 08 Jul)
Filter Analysis	7 July 03	-	9 Jul 03	1 Aug 03	Arista, 1400 samples analyzed
SAP	21 Jul 03		27 Jun 03		Working on
DB-Lock					
Clinical		18 Jul 03		21 Jul 03	CCSR to QA wk of 25 Aug 03, need edit checks
Analytical		08 Sep 03			Without MHBMA (MHBMA
Statistical		22 Sep 03			locks 9/29 + 2 weeks for tables/figures)
Preliminary Data					
Figs & Tabs		31 July 03 13 Oct 03		31 Jul 03	COHB and topography tables Analytical tables and figures
Monitoring	22 July 03	<u>Mid Sep</u>			22-25 July, 12-15 Aug, 19-22 Aug, 26-29 Aug
Draft Report		1 Dec 03			
Draft Report Review	1 Dec 03	15 Dec 03			
Final Report		31 Dec 03			

Issues/Decisions	Resolution
1- MHBMA analysis should remain in urinary biomarkers for study	13 Aug 03

Action items	Due Date	Done	MDS	PM
1. Summary tables for 3-HPMA data sent to PM as soon as possible	27 Aug 03	13 Aug 03	NW	MG
2. Analysis plan details for filter analysis provided to MDS	21 Aug 03	20 Aug 03	NW	QL

ACCORD SERIES K Short-term Study - EHCK6/01/03 (AA02981)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	-	-	23 Jan 03	30 Apr 03	
IRB Review	-	06 May 03	-	06 May 03	Approved (dated 01 May 03)
Amendments					
LOAC #1	-	-		27 May 03	IRB Approved (dated 22 May 03)
LOAC #2				17 Jun 03	IRB Approved (dated 17 Jun 03)-
					Approved 10 Jun (dated 09 Jun 03)
Nr1	06 Jun	10 Jun	06 Jun	10 Jun 03	
Recruiting	29 Apr 03	2 Jun 03	29 Apr 03	02 Jun 03	470 subjects as of 4 Jun
Screening	3 Jun 03	12 Jun 03	03 Jun 03		June 3-6 screen, June 11-12 phys. Additional screening 11 Jun cancelled, phys 12 Jun
Agreements/Proposals					
Recruitment (PA)	-	-	14 Apr 03	29 Apr 03	MDS #42, Advertising, ICF
Screening, execution (PA)	15 May 03	02 Jun 03	16 May 03	02 Jun 03	MDS #44
44 Amend 1 for Prot Amdmt 1	-	21 Jul 03	18 Jul 03	30 Jul 03	Sent to MDS 30 Jul 03
Analysis-Scherer (PA)	14 May 03	29 Jun 03	19 May 03		#7, valine adducts
Analysis-Arista (AP/PO)	14 May 03	29 Jun 03	14 May 03	27 May 03	filter analysis
Analysis-Covance (PA)	14 May 03	29 Jun 03	28 May 03	30 Jul 03	#14, caffeine, 8-epi, 11-dehy
Monitoring (AP/PO)	14 May 03	29 Jun 03	25 May 03	08 Jul 03	Clinimetrics
Study Conduct	19 Jun 03	30 Jun 03	19 Jun 03	30 June 03	120 Subjects randomized and completed
Topography Data	30 Jun 03	15 Jul 03	30 Jun 03	15 Jul 03	
Lab Analysis (clin and ext)					
COHb	04 Jun 03	15 Jul 03	04 Jun 03	15 Jul 03	MDS
4-ABP	<u>Needed</u>	<u>TBD</u>			Covance
LDL, HDL, Hem, etc	<u>Needed</u>	<u>TBD</u>			MDS
Fib, Von, HSC, Alb	<u>Needed</u>	<u>TBD</u>			Esoterix/ UNMC
Valines	7 July 03	9 Aug 03			ABF (shipped 08 Jul)
Filter/Analysis <i>N = 1409</i>	7 July 03	<u>TBD</u>	09 Jul 03	<u>08 Jul 03</u>	Arista (shipped 08 Jul) <i>DL to proper data folder</i>
SAP		21 Jul 03		<u>Needed</u>	<i>DL to proper data folder</i>
DB-Lock					
Clinical		18 Jul 03		<u>21 Jul 03</u>	
Analytical		<u>TBD</u>		<u>Needed</u>	
Statistical		<u>TBD</u>		<u>Needed</u>	Pending further Scheduling
Preliminary Data					
Figs & Tabs		31 July 03			<i>ES graphs sent to Prof. W</i>
Monitoring	22 July 03	15 Aug 03			22-25 July, 12-15 Aug
Draft Report		30 Nov 03			
Draft Report Review	1 Dec 03	15 Dec 03			
Final Report		31 Dec 03			

Issues/Decisions	Resolution
1. Confirm whether MHBMA should remain in urinary biomarkers for study	<i>7/23/03 do SAP, ASK 026/03</i>

Action items	Due Date	Done	MDS	PM
1. PA 44 amendment - MDS to Sign	19 Jun 03	18 Jul 03	KP	VK
2. Discussion around ETS analysis/process. Schedule teleconference.	23 Jul 03		KP	JO

50 sent to new guidance

new: do 21 2003 08 01
 signed today
 before in TACCORD SERIES K Short-term Study - EHCK6/01/03 (AA02981)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	-	-	23 Jan 03	30 Apr 03	
IRB Review	-	06 May 03	-	06 May 03	Approved (dated 01 May 03)
Amendments					
LOAC #1	-	-		27 May 03	IRB Approved (dated 22 May 03)
LOAC #2				17 Jun 03	IRB Approved (dated 17 Jun 03)
Nr1	06 Jun	10 Jun	06 Jun	10 Jun 03	Approved 10 Jun (dated 09 Jun 03)
Recruiting	29 Apr 03	2 Jun 03	29 Apr 03	02 Jun 03	470 subjects as of 4 Jun
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Agreements/Proposals					
Recruitment (PA)	-	-	14 Apr 03	29 Apr 03	MDS #42, Advertising, ICF
Screening, execution (PA)	15 May 03	02 Jun 03	16 May 03	02 Jun 03	MDS #44
44 Amend 1 for Prot Amend 1	-	21 Jul 03	18 Jul 03	30 Jul 03	To MDS 30 Jul 03-fully signed
Analysis-Scherer (PA)	14 May 03	29 Jun 03	19 May 03	08 Aug 03	#7, valine adducts
Analysis-Arista (AP/PO)	14 May 03	29 Jun 03	14 May 03	27 May 03	filter analysis
Analysis-Covance (PA)	14 May 03	29 Jun 03	28 May 03	30 Jul 03	#14, caffeine, 8-epi, 11-dehy
Monitoring (AP/PO)	14 May 03	29 Jun 03	25 May 03	08 Jul 03	Clinimetrics
Study Conduct	19 Jun 03	30 Jun 03	19 Jun 03	30 June 03	120 Subjects randomized and completed
Topography Data	30 Jun 03	15 Jul 03	30 Jun 03	15 Jul 03	
Lab Analysis (clin and ext)					
COHb	04 Jun 03	15 Jul 03	04 Jun 03	15 Jul 03	MDS
Caffeine Analysis				01 Aug 03	Covance
4-ABP	<u>Needed</u>	<u>TBD</u>			Covance
LDL, HDL, Hem, etc	<u>Needed</u>	<u>TBD</u>			MDS
Fib, Von, HSC, Alb	<u>Needed</u>	<u>TBD</u>			Esoterix/ UNMC
Valines	7 July 03	31 Sep 03	09 Jul 03		ABF (shipped 08 Jul)
Filter Analysis	7 July 03	-	9 Jul 03	1 Aug 03	Arista, 1400 samples analyzed
SAP	21 Jul 03		27 Jun 03		Working on
DB-Lock					
Clinical		18 Jul 03		21 Jul 03	
Analytical		08 Sep 03			Without MHBMA (MHBMA locks 9/29 + 2 weeks for tables/figures)
Statistical		22 Sep 03			
Preliminary Data					
Figs & Tabs		31 July 03 13 Oct 03		31 Jul 03	COHB and topography tables Analytical tables and figures
Monitoring	22 July 03	22 Aug 03		~ 15 Sep	22-25 July, 12-15 Aug, 19-22 Aug
Draft Report		30 Nov 03			
Draft Report Review	1 Dec 03	15 Dec 03			
Final Report		31 Dec 03			

Issues/Decisions	Resolution
1. MHBMA analysis should remain in urinary biomarkers for study	13 Aug 03

Action items	Due Date	Done	MDS	PM
1. Discussion around ETS analysis/process. Schedule teleconference. ETS data sent to Nancy week of 07/28/03.	23 Jul 03	13 Aug 03	KP	JO
2. Analysis plan details for filter analysis provided to MDS	21 Aug 03		NW	QL

SAP p. 96, 100 → all

ACCORD SERIES K Long-term Study - EHCK6/02/03 (MDS #AA03468)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	TBD	30 Sep 03	08 Aug 03		To send draft by 08 Sep to meet timelines
IRB Review	1 Oct 03	7 Oct 03			
Amendments Nr. 1					
Recruiting	TBD	TBD			
Screening	TBD	TBD			
Agreements/Proposals					
Study Conduct	13 Nov 03	29 April 04			Target 100 subjects. Prefer to use as many of the subjects from the EHCK6/01/03 study as possible.
Topography Data					<i>but failed to query a PC SD</i>
Lab Analysis (clin and ext)					<i>soft 0?? -- will call MDS</i>
SAP					
DB-Lock					
Clinical		13 May 04			
Analytical		12 Aug 04			
Statistical		26 Aug 04			
Preliminary Data					
Monitoring					
Draft Report		21 Oct 04			
Draft Report Review		04 Nov 04			
Final Report		02 Dec 04			

Issues/Decisions	Resolution
1. Plan to use WatchPC device during study (to track daily cigarette consumption)	8 Aug 03
2.	

Action items	Due Date	Done	MDS	PM
1. Study confirmation, including recruiting/screening/study conduct dates	22 Aug 03		KP	VK
2. Postponement costs for Jun 03 needed	18 Aug 03		KP	VK
3. Estimate of subjects willing to continue from EHCK6/01/03	21 Aug 03		KP	MG
4. Decision on whether smoking at days -1 and day 8 need to be controlled (as in EHCK6/01/03)	21 Aug 03		MS	MG/BZ

Amcote (supp time 7th-11th)
?

if sep rand orders

ACCORD SERIES K Long-term Study - EHCK6/02/03 (MDS #TBD)

	Planned		Actual		Comments
	Start	Finish	Start	Finish	
Draft Protocol	TBD	20 Sep 03			<i>Mike working on final timeline</i>
IRB Review	1 Oct 03	7 Oct 03			
Amendments Nr. 1					
Recruiting	TBD	TBD			
Screening	TBD	TBD			
Agreements/Proposals	-	-			
Study Conduct	13 Nov 03	4/29/04			Target 100 subjects. Prefer to use as many of the subjects from the EHCK6/01/03 study as possible.
Topography Data					
Lab Analysis (clin and ext)					
SAP					
DB-Lock					
Preliminary Data					
Monitoring					
Draft Report					
Draft Report Review					
Final Report					

Issues/Decisions	Resolution
1.	
2.	

Action items	Due Date	Done	MDS	PM
1. Study outline	1 Aug 03		KP	MG
2. Study confirmation, including recruiting/screening/study conduct dates	1 Aug 03		KP	VK